

## EAST Search History

| Ref # | Hits | Search Query   | DBs                | Default Operator | Plurals | Time Stamp       |
|-------|------|--|--------------------|------------------|---------|------------------|
| L1    | 882  | 514/375.ccls.  | US-PGPUB;<br>USPAT | OR               | ON      | 2007/09/19 14:06 |
| L2    | 1497 | 514/381.ccls.  | US-PGPUB;<br>USPAT | OR               | ON      | 2007/09/19 14:06 |
| L3    | 1128 | 514/411.ccls.  | US-PGPUB;<br>USPAT | OR               | ON      | 2007/09/19 14:06 |
| L4    | 605  | 514/418.ccls.  | US-PGPUB;<br>USPAT | OR               | ON      | 2007/09/19 14:06 |
| L5    | 1652 | 514/419.ccls.  | US-PGPUB;<br>USPAT | OR               | ON      | 2007/09/19 14:06 |
| L6    | 232  | 514/429.ccls.  | US-PGPUB;<br>USPAT | OR               | ON      | 2007/09/19 14:06 |
| L7    | 271  | 514/431.ccls.  | US-PGPUB;<br>USPAT | OR               | ON      | 2007/09/19 14:07 |
| L8    | 417  | 514/448.ccls.  | US-PGPUB;<br>USPAT | OR               | ON      | 2007/09/19 14:07 |
| L9    | 792  | 514/569.ccls.  | US-PGPUB;<br>USPAT | OR               | ON      | 2007/09/19 14:07 |
| L10   | 1111 | 514/570.ccls.  | US-PGPUB;<br>USPAT | OR               | ON      | 2007/09/19 14:07 |
| L11   | 7750 | L1 OR L2 OR L3 OR L4 OR L5 OR<br>L6 OR L7 OR L8 OR L9 OR L10 | US-PGPUB;<br>USPAT | OR               | ON      | 2007/09/19 14:07 |
| L12   | 1526 | L11 AND ALZHEIMER  | US-PGPUB;<br>USPAT | OR               | ON      | 2007/09/19 14:08 |
| L13   | 265  | L12 AND AMYLOID  | US-PGPUB;<br>USPAT | OR               | ON      | 2007/09/19 14:08 |

# STN Search (Reg/Caplus) - Claim 42

10/540, 601

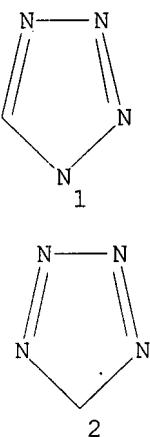
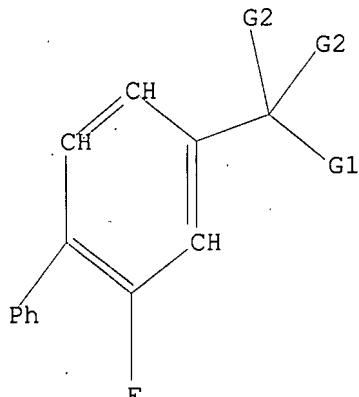
09/19/2007

L1 STRUCTURE UPLOADED

=> D

L1 HAS NO ANSWERS

L1 STR



G1 CO2H, COOH, [@1], [@2]

G2 Cb, Ak

Structure attributes must be viewed using STN Express query preparation.

=> S L1

SAMPLE SEARCH INITIATED 14:57:13 FILE 'REGISTRY'  
SAMPLE SCREEN SEARCH COMPLETED - 19 TO ITERATE

100.0% PROCESSED 19 ITERATIONS 0 ANSWERS  
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*  
BATCH \*\*COMPLETE\*\*

PROJECTED ITERATIONS: 119 TO 641  
PROJECTED ANSWERS: 0 TO 0

L2 0 SEA SSS SAM L1

=> S L1 FULL  
FULL SEARCH INITIATED 14:57:18 FILE 'REGISTRY'  
FULL SCREEN SEARCH COMPLETED 455 TO ITERATE

100.0% PROCESSED ✓ 455 ITERATIONS  
SEARCH TIME: 00.00.01

7 ANSWERS

L3 7 SEA SSS FUL L1

=> FIL CAPLUS

10/540,601

09/19/2007

| COST IN U.S. DOLLARS | SINCE FILE ENTRY | TOTAL SESSION |
|----------------------|------------------|---------------|
| FULL ESTIMATED COST  | 172.10           | 172.31        |

FILE 'CAPLUS' ENTERED AT 14:57:24 ON 19 SEP 2007  
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FILE LAST UPDATED: 18 Sep 2007 (20070918/ED)

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=> S L3  
L4            23 L3

=> S L4 AND ALZHEIMER  
        46782 ALZHEIMER  
L5            8 L4 AND ALZHEIMER

=> D IBIB ABS HITSTR 1-8

✓

L5 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2007:384840 CAPLUS

DOCUMENT NUMBER: 146:365795

TITLE: Combination therapy comprising Aβ42 lowering agent and hormonal modulating agent

INVENTOR(S): Hobden, Adrian; Leslie, Wayne

PATENT ASSIGNEE(S): Myriad Genetics, Incorporated, USA

SOURCE: U.S. Pat. Appl. Publ., 24pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO.             | KIND | DATE     | APPLICATION NO. | DATE       |
|------------------------|------|----------|-----------------|------------|
| US 2007078114          | A1   | 20070405 | US 2006-470190  | 20060905   |
| PRIORITY APPLN. INFO.: |      |          | US 2005-713369P | P 20050902 |

OTHER SOURCE(S): MARPAT 146:365795

AB The invention relates to combinations of one or more Aβ42 lowering agents and one or more hormonal modulating agent for treatment of Alzheimer's disease and other diseases. Thus, tablet was prepared containing (R)-2-(2-Fluoro-4-biphenyl)propionic acid 400 mg,

microcryst. cellulose 392 mg, colloidal silicon dioxide 4 mg, magnesium stearate 4 mg, and prednisone 25 mg.

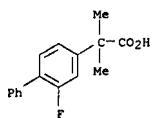
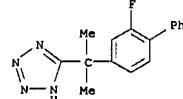
IT 150972-50-2 730977-79-4 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (combination therapy comprising Aβ42 lowering agent and hormonal modulating agent)

RN 150972-50-2 CAPLUS

CN [1,1'-Biphenyl]-4-acetic acid, 2-fluoro-a,a-dimethyl- (CA INDEX NAME)

L5 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

(Continued)

RN 730977-79-4 CAPLUS  
CN 2H-Tetrazole, 5-[1-(2-fluoro[1,1'-biphenyl]-4-yl)-1-methylethyl]- (CA INDEX NAME)

L5 ANSWER 2 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2007:63637 CAPLUS

DOCUMENT NUMBER: 146:135610

TITLE: Methods of treating overactive bladder and urinary incontinence

INVENTOR(S): Laughlin, Mark

PATENT ASSIGNEE(S): Myriad Genetics, Incorporated, USA

SOURCE: U.S. Pat. Appl. Publ., 16pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO.             | KIND | DATE     | APPLICATION NO. | DATE       |
|------------------------|------|----------|-----------------|------------|
| US 2007015832          | A1   | 20070118 | US 2006-487177  | 20060714   |
| PRIORITY APPLN. INFO.: |      |          | US 2005-699727P | P 20050714 |

OTHER SOURCE(S): MARPAT 146:135610

AB The invention relates to methods of treating or slowing the onset of overactive bladder and urinary incontinence, or a symptom thereof selected

from urinary frequency, urinary urgency, nocturia, or enuresis. In one aspect, the method comprises identifying and administering to a subject

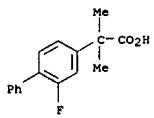
in need of treatment a therapeutically effective amount of an Aβ42 lowering agent such as (R)-2-(2-fluoro-4-biphenyl)propionic acid.

IT 150972-50-2 730977-79-4 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(methods of treating overactive bladder and urinary incontinence using Aβ42 lowering agents such as (fluorobiphenyl)propionic acid and combination with other agents in relation to Alzheimer's disease)

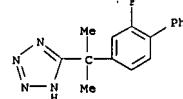
RN 150972-50-2 CAPLUS

CN [1,1'-Biphenyl]-4-acetic acid, 2-fluoro-a,a-dimethyl- (CA INDEX NAME)

RN 730977-79-4 CAPLUS  
CN 2H-Tetrazole, 5-[1-(2-fluoro[1,1'-biphenyl]-4-yl)-1-methylethyl]- (CA INDEX NAME)

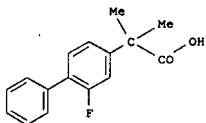
L5 ANSWER 2 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

(Continued)



L5 ANSWER 3 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN  
 ACCESSION NUMBER: 2006:232875 CAPLUS  
 DOCUMENT NUMBER: 144:403783

TITLE: The geminal dimethyl analogue of Flurbiprofen as a novel Aβ42 inhibitor and potential Alzheimer's disease modifying agent  
 AUTHOR(S): Stock, Nicholas; Munoz, Benito; Wrigley, Jonathan D. J.; Shearman, Mark S.; Behar, Dirk; Peachey, James; Williamson, Toni L.; Bain, Gretchen; Chen, Weichao; Jiang, Xiaohui; St-Jacques, Rene; Prasit, Peppe Department of Chemistry, Merck Research Laboratories, San Diego, CA, 92121, USA  
 CORPORATE SOURCE: Bioorganic & Medicinal Chemistry Letters (2006), 16(8), 2219-2223  
 SOURCE: CODEN: BMCLB6; ISSN: 0960-894X  
 PUBLISHER: Elsevier B.V.  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 GI



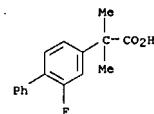
I

AB The subtle modification of a selection of Aβ42 inhibiting non-steroidal anti-inflammatory drugs (NSAIDs), through synthesis of the geminal di-Me analogs, was anticipated to ablate their cyclooxygenase activity while maintaining Aβ42 inhibition. Methylflurbiprofen I exhibited similar in vitro Aβ42 inhibition to its parent NSAID Flurbiprofen and was further evaluated in the Tg2576 mouse model of Alzheimer's disease and an animal model of gastro-intestinal (GI) impairment, but proved unviable for further clin. development.

IT 150972-50-2P  
 RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 . (geminal di-Me analog of Flurbiprofen as Aβ42 inhibitor and potential Alzheimer's disease modifying agent)

RN 150972-50-2 CAPLUS  
 CN (1,1'-Biphenyl)-4-acetic acid, 2-fluoro-a,a-dimethyl- (CA INDEX NAME)

L5 ANSWER 3 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)



REFERENCE COUNT: 35 THERE ARE 35 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN  
 ACCESSION NUMBER: 2006:167837 CAPLUS  
 DOCUMENT NUMBER: 144:239971

TITLE: Pharmaceutical composition and method for treating neurodegenerative disorders

INVENTOR(S): Hobden, Adrian  
 PATENT ASSIGNEE(S): Myriad Genetics, Inc., USA  
 SOURCE: PCT Int'l Appl., 64 pp.

DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

| PATENT NO.    | KIND   | DATE     | APPLICATION NO. | DATE     |
|---------------|--|----------|-----------------|----------|
| WO 2006020850 | A2   | 20060223 | WO 2005-US28714 | 20050811 |
| WO 2006020850 | A3   | 20060504 |                 |          |
| W:            | AF, AG, AL, AM, AT, AU, A2, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SI, SM, SR, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW |          |                 |          |
| RW:           | AT, BE, BG, CH, CT, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM   |          |                 |          |

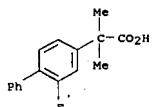
PRIORITY APPLN. INFO.: US 2004-600447P P 20040811

AB The invention provides compns. and methods for treating neurodegenerative disorders. The method of the invention involves administering to an individual in need of treatment a composition having an acetylcholine esterase

inhibitor and another therapeutic agent. The methods and compns. of the invention are useful for treating and preventing neurodegenerative disorders like Alzheimer's disease, dementia, and mild cognitive impairment.

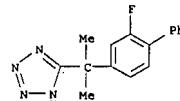
IT 150972-50-2 730977-79-4  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (pharmaceutical composition and method for treating neurodegenerative disorders)

RN 150972-50-2 CAPLUS  
 CN (1,1'-Biphenyl)-4-acetic acid, 2-fluoro-a,a-dimethyl- (CA INDEX NAME)



L5 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)

RN 730977-79-4 CAPLUS  
 CN 2H-Tetrazole, 5-[1-(2-fluoro[1,1'-biphenyl]-4-yl)-1-methylethyl]- (CA INDEX NAME)



L5 ANSWER 5 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2006:164779 CAPLUS

DOCUMENT NUMBER: 144:239954

TITLE: Pharmaceutical compositions acetylcholine esterase

inhibitors for treating neurodegenerative disorders

INVENTOR(S): Hobden, Adrian

PATENT ASSIGNEE(S): Myriad Genetics, Inc., USA

SOURCE: PCT Int. Appl., 64 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO.    | KIND | DATE     | APPLICATION NO. | DATE     |
|---------------|------|----------|-----------------|----------|
| WO 2006020852 | A2   | 20060223 | WO 2005-US28716 | 20050811 |
| WO 2006020852 | A3   | 20060526 |                 |          |

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW  
RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

PRIORITY APPN. INFO.: US 2004-600600P P 20040811

AB The invention provides compns. and methods for treating neurodegenerative disorders. The method of the invention involves administering to an individual in need of treatment an acetylcholine esterase inhibitor in combination with another therapeutic agent. The methods and compns. of the invention are useful for treating and preventing neurodegenerative disorders like Alzheimer's disease, dementia, and mild cognitive impairment. A tablet contained (R)-2-(2-fluoro-4-biphenyl)propionic acid 400, microcryst. cellulose 392, colloidal silicon dioxide 4, magnesium stearate 4, and galantamine hydrobromide 16 mg.

IT 150972-50-2 730977-79-4

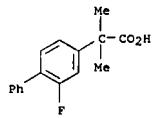
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(pharmaceutical compns. acetylcholine esterase inhibitors for treating neurodegenerative disorders)

RN 150972-50-2 CAPLUS

CN [1,1'-Biphenyl]-4-acetic acid, 2-fluoro- $\alpha,\alpha$ -dimethyl- (CA INDEX NAME)

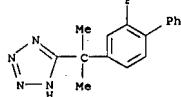
L5 ANSWER 5 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

(Continued)



RN 730977-79-4 CAPLUS

CN 2H-Tetrazole, 5-(1-(2-fluoro-4-biphenyl)-4-yl)-1-methylethyl- (CA INDEX NAME)



L5 ANSWER 6 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2006:164727 CAPLUS

DOCUMENT NUMBER: 144:260786

TITLE: Pharmaceutical compositions containing acetylcholine esterase inhibitors for treating neurodegenerative disorders

INVENTOR(S): Hobden, Adrian

PATENT ASSIGNEE(S): Myriad Genetics, Inc., USA

SOURCE: PCT Int. Appl., 64 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO.    | KIND | DATE     | APPLICATION NO. | DATE     |
|---------------|------|----------|-----------------|----------|
| WO 2006020853 | A2   | 20060223 | WO 2005-US28717 | 20050811 |
| WO 2006020853 | A3   | 20060526 |                 |          |

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW  
RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

PRIORITY APPN. INFO.: US 2004-600470P P 20040811

AB The invention provides compns. and methods for treating neurodegenerative disorders. The method of the invention involves administering to an individual in need of treatment a composition having an acetylcholine esterase

inhibitor and another therapeutic agent. The methods and compns. of the invention are useful for treating and preventing neurodegenerative disorders like Alzheimer's disease, dementia, and mild cognitive impairment. A tablet contained (R)-2-(2-fluoro-4-biphenyl)propionic acid 400, microcryst. cellulose 392, colloidal silicon dioxide 4, magnesium stearate 4, and donepezil hydrochloride 16 mg.

IT 150972-50-2

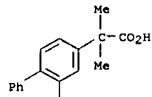
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(pharmaceutical composition and method for treating neurodegenerative disorders)

RN 150972-50-2 CAPLUS

CN [1,1'-Biphenyl]-4-acetic acid, 2-fluoro- $\alpha,\alpha$ -dimethyl- (CA INDEX NAME)

L5 ANSWER 6 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

(Continued)



L5 ANSWER 7 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2005:1075559 CAPLUS

DOCUMENT NUMBER: 143:367205

TITLE: Preparation of compounds, especially indoles and biphenyls, useful for treating neurodegenerative disorders, particularly Alzheimer's disease and other amyloid B42 protein-related disorders

INVENTOR(S): Slade, Rachel M.; Weiner, Warren S.; Delmar, Eric G.; Klimova, Yevgeniya I.; Trovato, Richard

PATENT ASSIGNEE(S): Myriad Genetics, Inc., USA

SOURCE: PCT Int. Appl., 110 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO.    | KIND | DATE     | APPLICATION NO. | DATE     |
|---------------|------|----------|-----------------|----------|
| WO 2005092062 | A2   | 20051006 | WO 2005-US9595  | 20050321 |
| WO 2005092062 | A3   | 20060803 |                 |          |

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM,

ZW

RW: BW, GH, GM, KE, LS, MM, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, C2, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CH, GA, GN, GG, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.: US 2004-554571P P 20040319

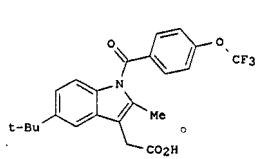
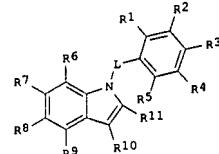
US 2004-590259P P 20040722

OTHER SOURCE(S): CASREACT 143:367205; MARPAT 143:367205

GI

L5 ANSWER 7 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

(Continued)



AB Title compd., e.g. I, [L = C:O, CH2; R1, R2, R4-R7, R9 = independently OH, halo, haloalkyl, etc.; R3 = CHF2, CF3, OCF3, OCHF2; R8 = H, halo, alkoxy, etc.; R10 = alkylene-CO2H, alkylene-CO2H, alkynylene-CO2H; R11 = alkyl; and their pharmaceutically acceptable salts] were prepared as lowering cellular amyloid B42 protein (Aβ42) production and/or secretion agents useful for the therapeutic treatment and prevention of neurodegenerative disorders such as Alzheimer's disease, mild cognitive impairment, dementia etc. For example, II was prepared in 3 steps

from [5-(tert-butyl)-2-methyl-1H-indol-3-yl]acetic acid and 4-trifluoromethoxybenzoyl chloride. Selected I were found to not significantly inhibit COX-1 and COX-2 at 100 μM. In amyloid precursor protein assays selected I lowered Aβ42 levels by at least 50% of DMSO control at concentration ranging from 30 to 80 μM. IT 150972-50-2P, 2-(2-Fluorobiphenyl-4-yl)-2-methylpropanoic acid 866235-88-3P, 2-(2-Fluorobiphenyl-4-yl)-2-methylbutyric acid 866235-92-9P, 2-Ethyl-2-(2-fluorobiphenyl-4-yl)butyric acid (Uses) PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

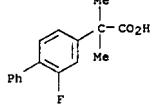
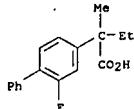
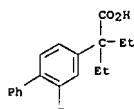
(drug candidate; preparation of compds., especially indoles and biphenyls, useful for treating neurodegenerative disorders)

RN 150972-50-2 CAPLUS

CN [1,1'-Biphenyl]-4-acetic acid, 2-fluoro- $\alpha$ , $\alpha$ -dimethyl- (CA

L5 ANSWER 7 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)

INDEX NAME)

RN 866235-88-3 CAPLUS  
CN [1,1'-Biphenyl]-4-acetic acid,  $\alpha$ -ethyl-2-fluoro- $\alpha$ -methyl- (CA INDEX NAME)RN 866235-92-9 CAPLUS  
CN [1,1'-Biphenyl]-4-acetic acid,  $\alpha$ , $\alpha$ -diethyl-2-fluoro- (CA INDEX NAME)

L5 ANSWER 8 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2004:633474 CAPLUS

DOCUMENT NUMBER: 141:162386

TITLE: Anti-Alzheimer compositions containing

geminally di-substituted NSAID derivatives

INVENTOR(S): Munoz, Benito; Prasit, Petpiboon; Stock, Nicholas

Simon

PATENT ASSIGNEE(S): Merck &amp; Co., Inc., USA

SOURCE: PCT Int. Appl., 40 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO.    | KIND | DATE     | APPLICATION NO. | DATE     |
|---------------|------|----------|-----------------|----------|
| WO 2004064771 | A2   | 20040805 | WO 2004-US424   | 20040109 |
| WO 2004064771 | A3   | 20041223 |                 |          |

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, AU 2004206796 A1 20040805 AU 2004-206796 20040109

CA 2512704 A1 20040805 CA 2004-2512704 20040109

EP 1587798 A2 20051026 EP 2004-701220 20040109

R: AT, BE, CH, DE, DK, ES, FR, GB, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK

JP 2006517925 T 20060803 JP 2006-500855 20040109

US 2006063937 A1 20060323 US 2005-540601 20050623

PRIORITY APPLN. INFO.: US 2003-439947P P 20030114

US 2003-439965P P 20030114

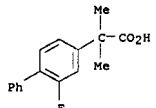
WO 2004-US424 W 20040109

OTHER SOURCE(S): MARPAT 141:162386

GI

AB The present invention encompasses compds. of Formula I or pharmaceutically acceptable salts thereof, wherein A is the base mol. of a propionic acid or acetic acid NSAID, or a derivative thereof; X is -CO2H, 1H-tetrazol-5-yl or 2H-tetrazol-5-yl and R1 and R2 are each independently selected from the group consisting of C1-6 alkyl and C3-6 cycloalkyl; R3 is independently selected from fluoro, chloro, bromo, NH2, Me, Et, methoxy and CF3. This invention also relates to the pharmaceutical composition comprising said compds. and methods of using said compds. The compds. of the present invention lower the level of Aβ42 and are therefore useful for

L5 ANSWER 8 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)  
 preventing, delaying or reversing the progression of Alzheimer's  
 Disease.  
 IT 150972-50-2 730977-79-4P  
 RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological  
 study); PREP (Preparation); USES (Uses)  
 (prep of geminally di-substituted NSAID derivs. as anti-  
 Alzheimer agents)  
 RN 150972-50-2 CAPLUS  
 CN [1,1'-Biphenyl]-4-acetic acid, 2-fluoro-a,a-dimethyl- (CA  
 INDEX NAME)



RN 730977-79-4 CAPLUS  
 CN 2H-Tetrazole, 5-[1-(2-fluoro[1,1'-biphenyl]-4-yl)-1-methylethyl]- (CA  
 INDEX NAME)

